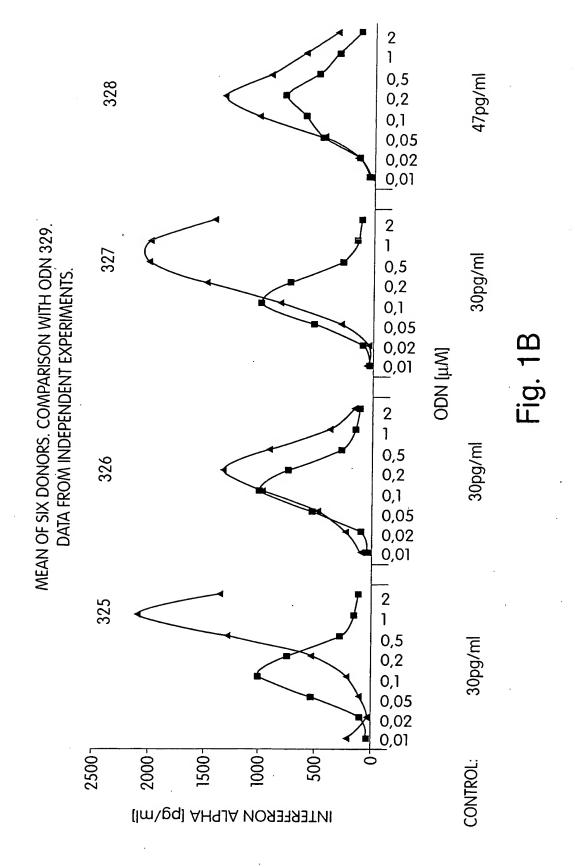
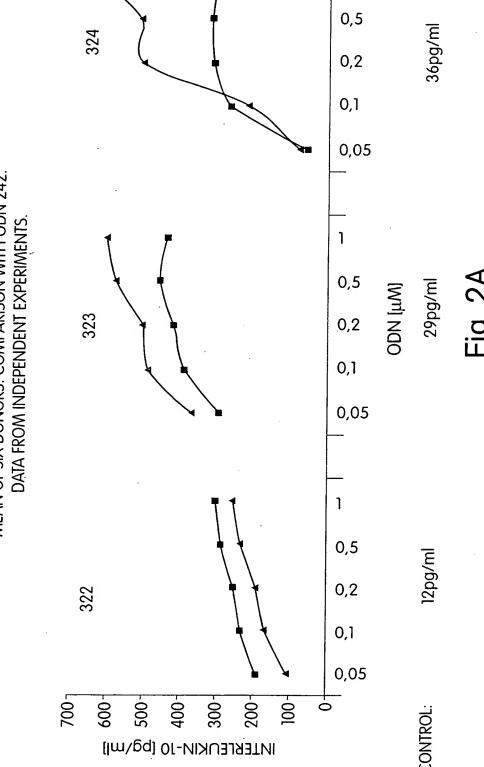


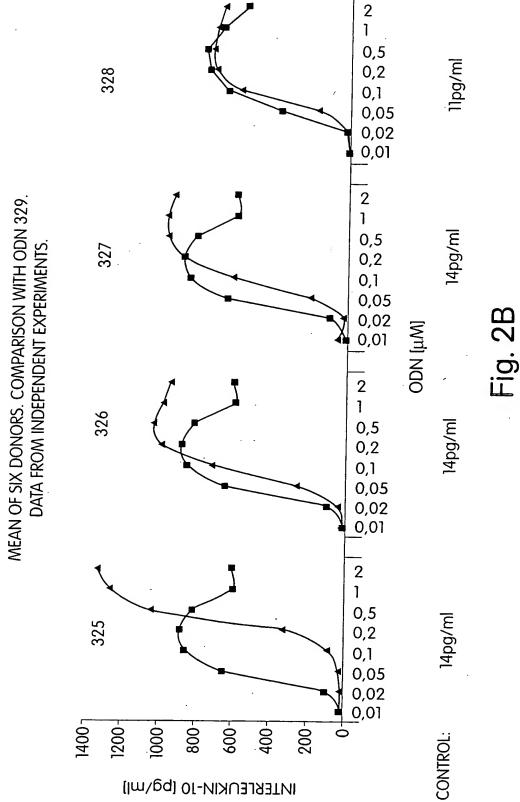
BEST AVAILABLE COPY





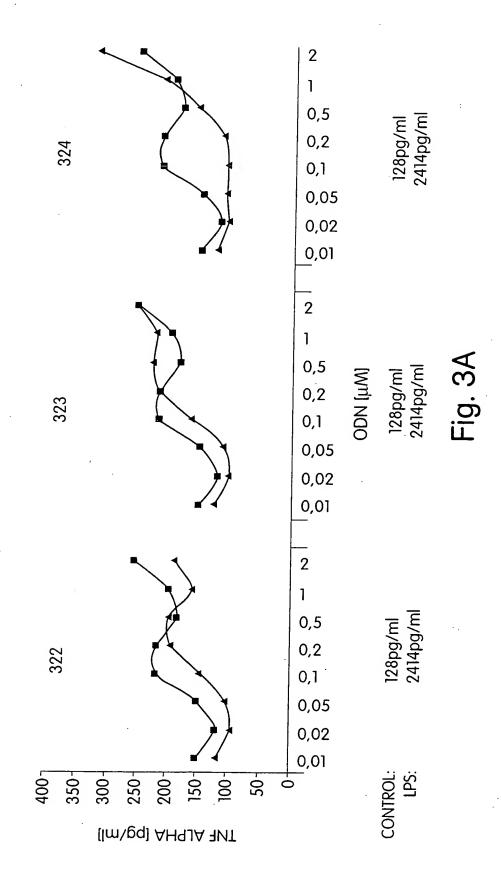
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MEAN OF SIX DONORS. COMPARISON WITH ODN 242.



Docket No.: C01037.70048.US

MEAN OF THREE DONORS. COMPARISON WITH ODN 329.



2 1 219pg/ml 1631pg/ml 0,5 0,2 328 0,1 0,05 0,02 <u>0</u>,01 2 MEAN OF THREE DONORS. COMPARISON WITH ODN 329. 1 128pg/ml 2414pg/ml 0,5 0,2 DATA FROM INDEPENDENT EXPERIMENTS. 327 0,1 0,05 0,02 0,01 [mrl] NGO 2 . 128pg/ml 2414pg/ml 1 0,5 0,2 326 0,1 0,05 0,02 0,01 2 1 128pg/ml 2414pg/ml 0,5 0,2 0,1 0,05 325 0,02 0,01 CONTRÓL: LPS: 100

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Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US

**600**1

500-

400-

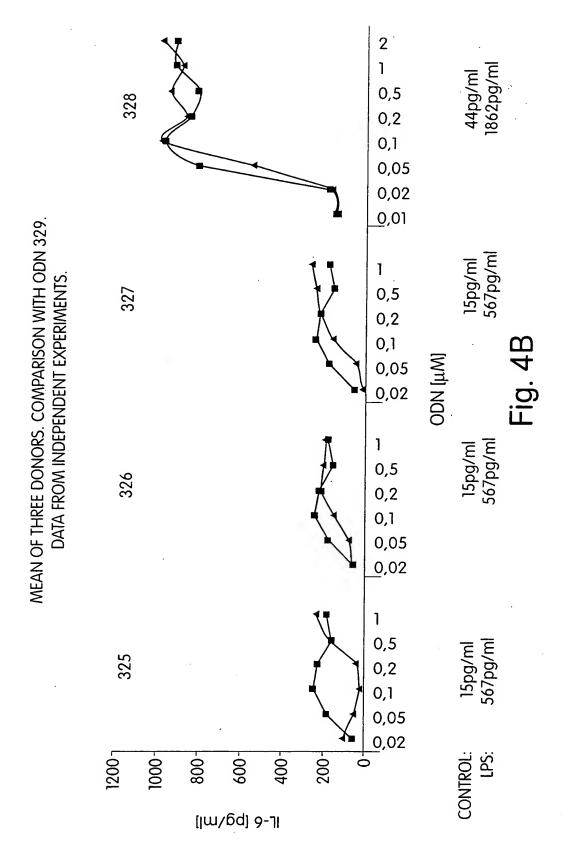
300-

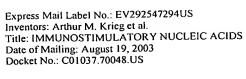
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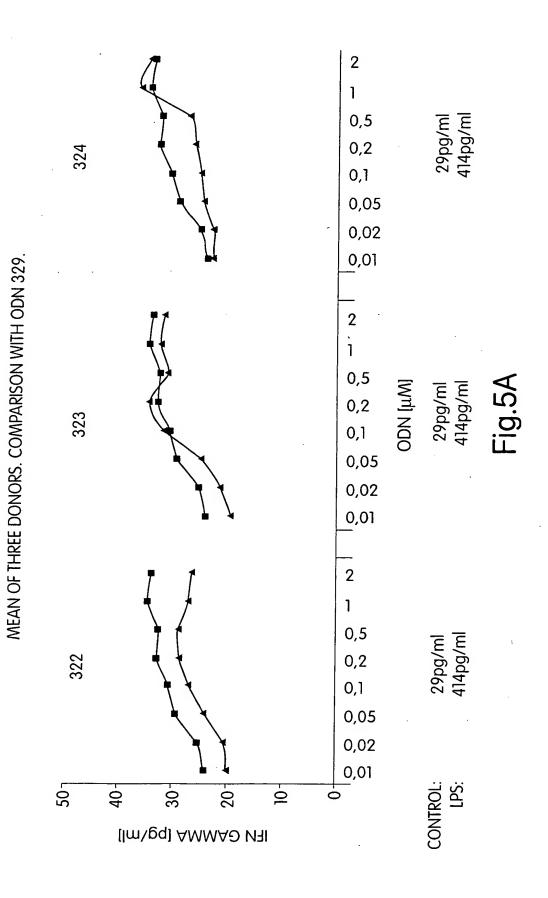
200-

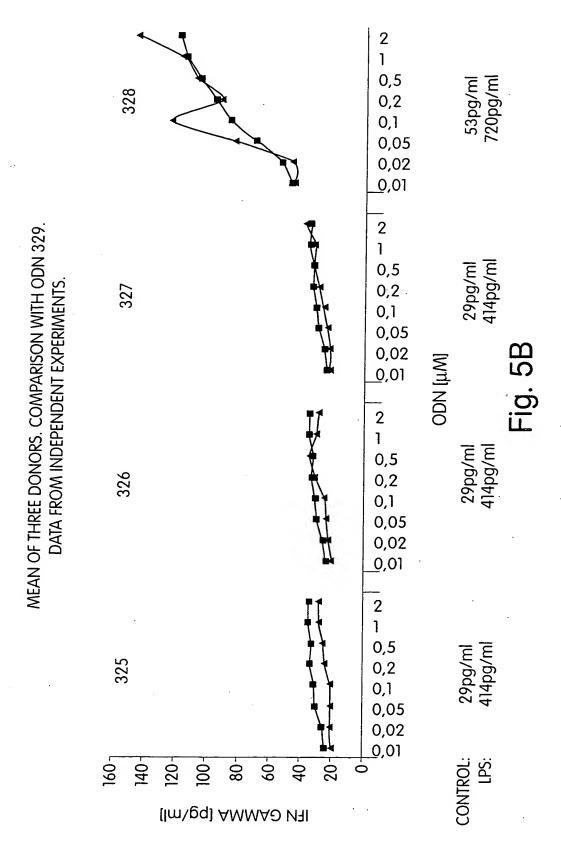
0,5 0,2 324 0,1 0,05 0,02 MEAN OF THREE DONORS. COMPARISON WITH ODN 329. DATA FROM INDEPENDENT EXPERIMENTS. 1 0,5 15pg/ml 567pg/ml (Wri) NGO 0,2 323 0,1 0,05 0,02 1 0,5 15pg/ml 567pg/ml 0,2 322 0,1 0,05 0,02 CONTROL: LPS: 500<sub>1</sub> 100 400-300-200-

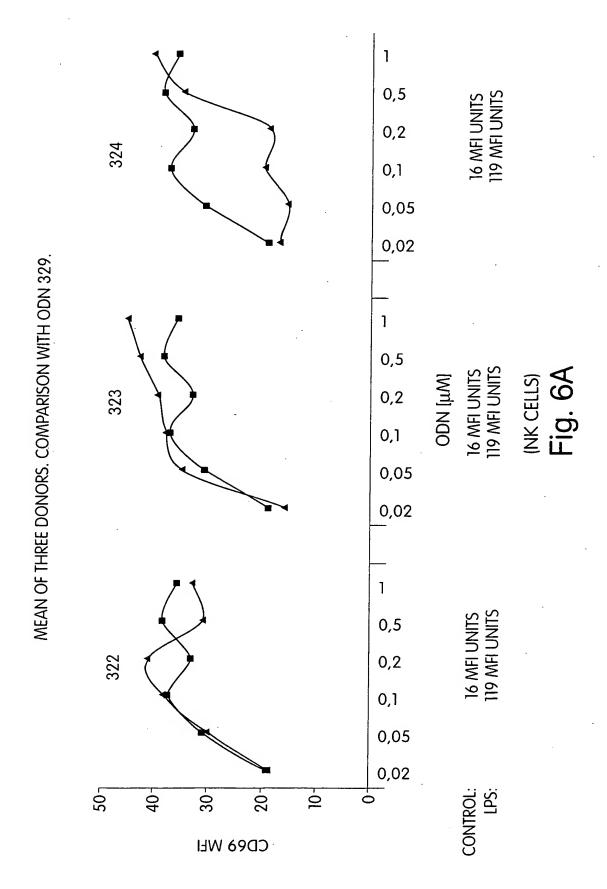
[w/6d] 9-71

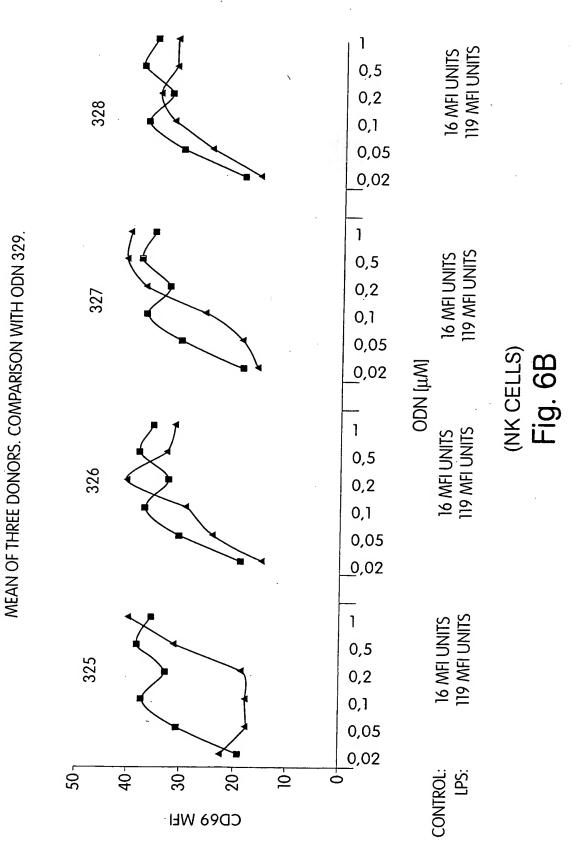


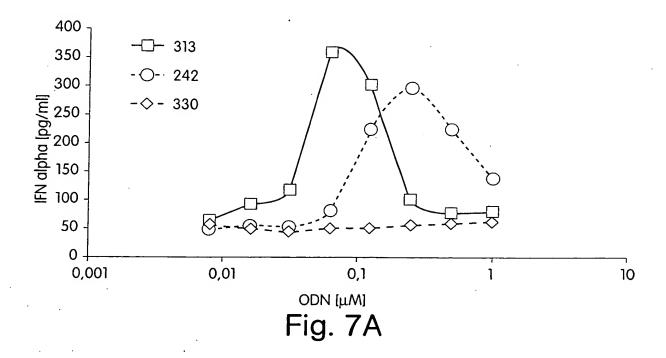


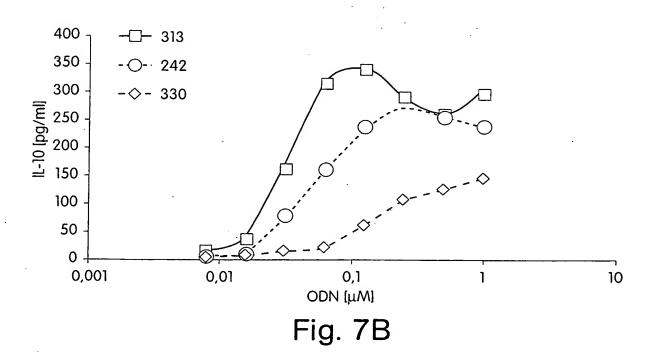


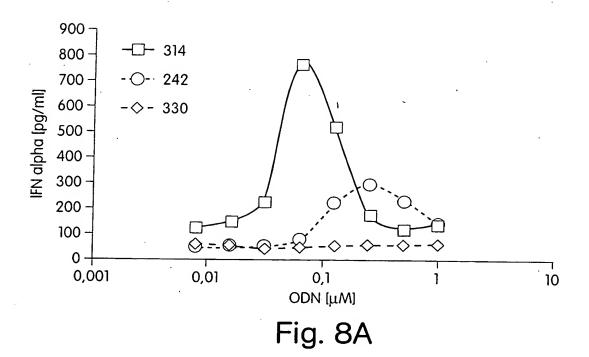


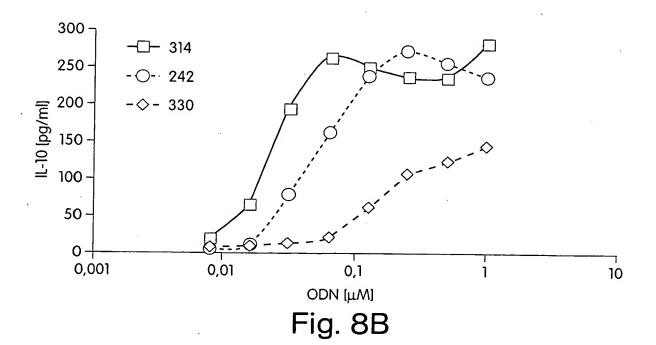


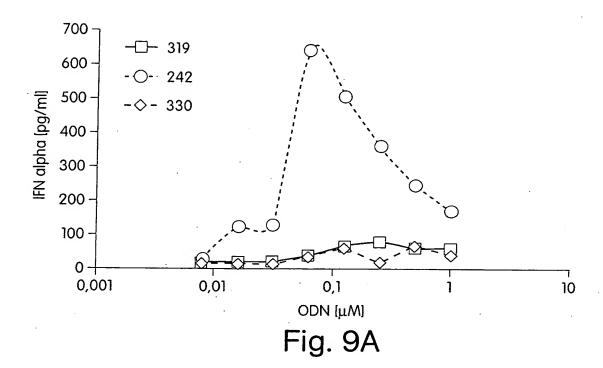


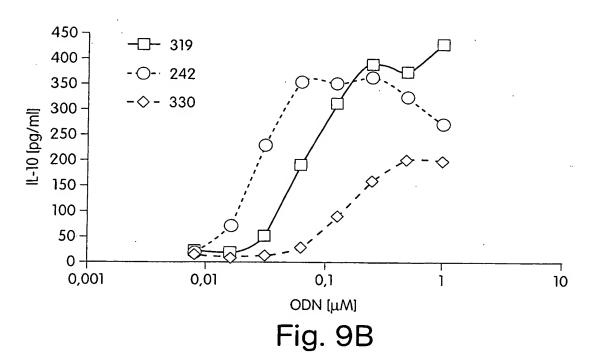












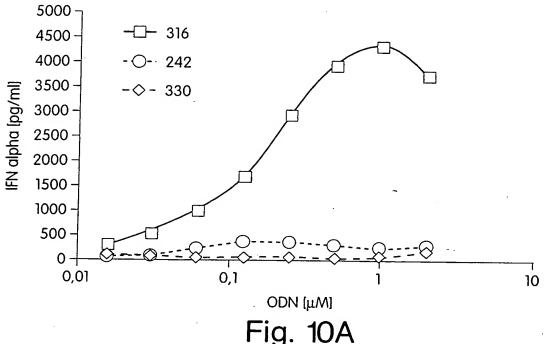


Fig. 10A

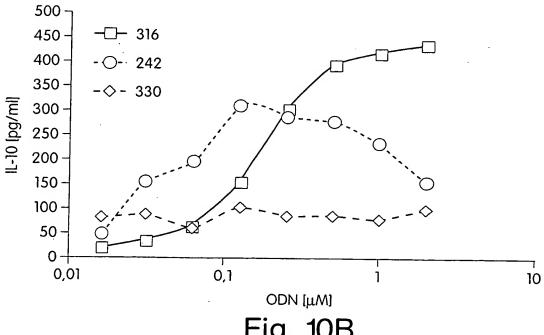
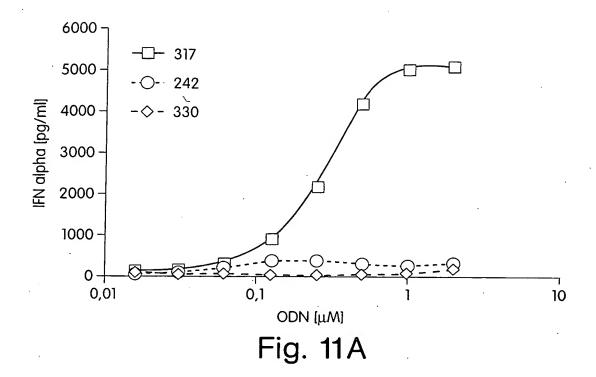
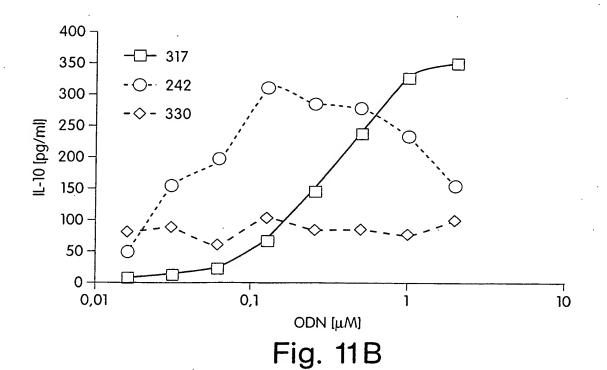
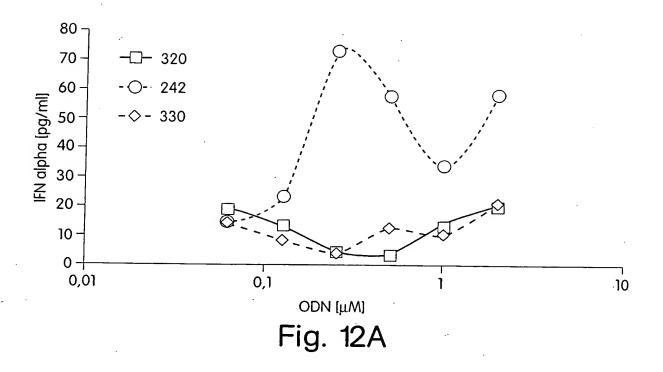
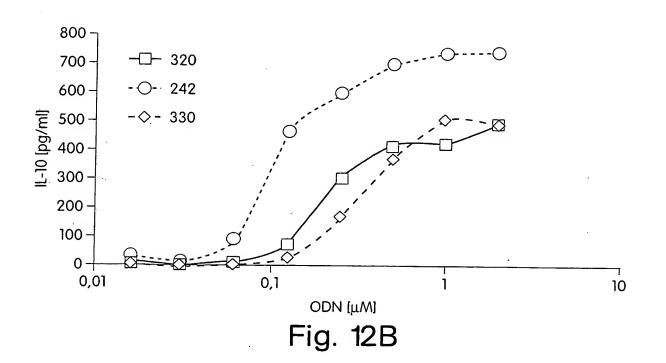


Fig. 10B

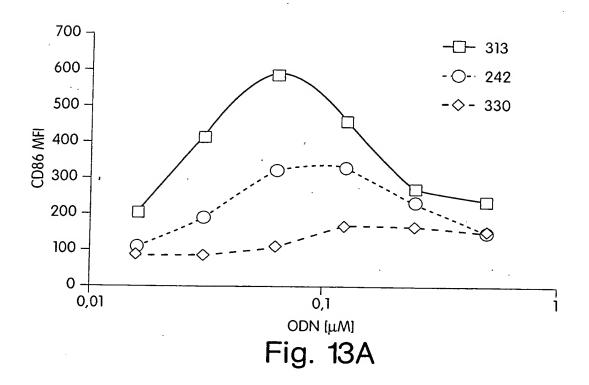


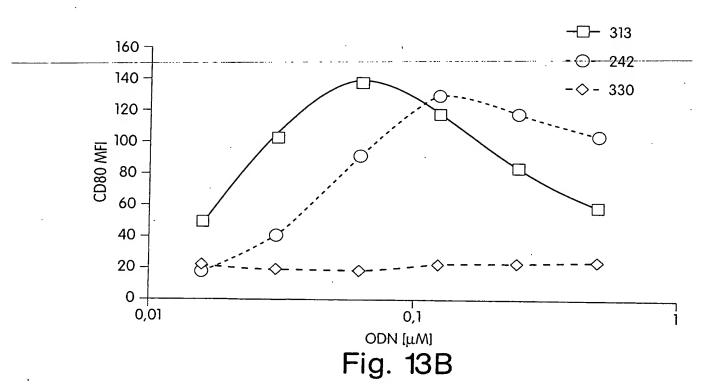






Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US





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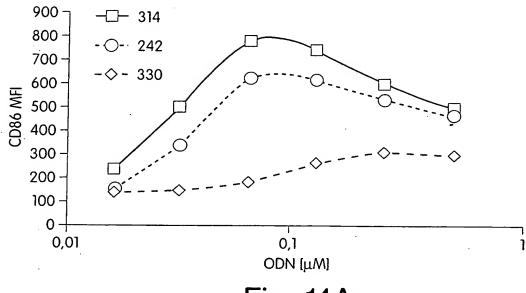


Fig. 14A

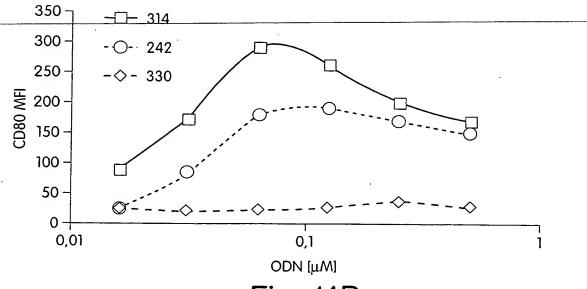
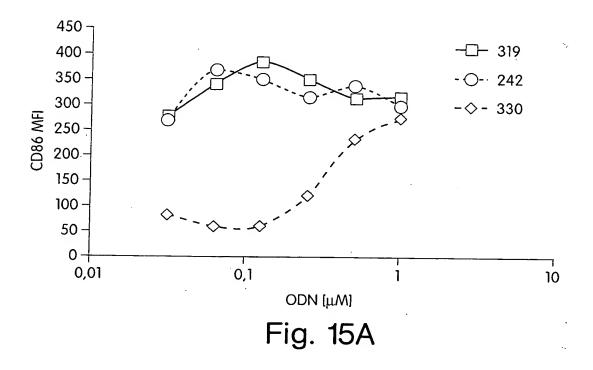
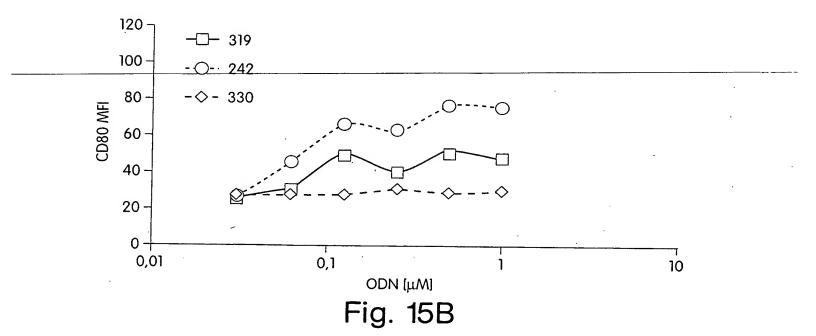
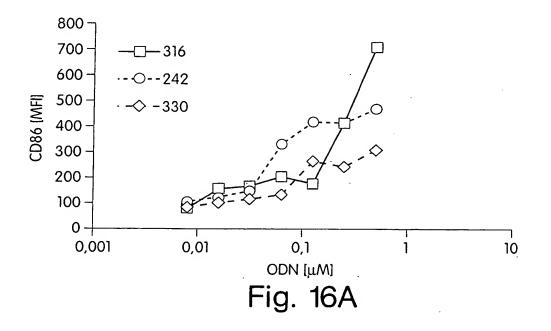
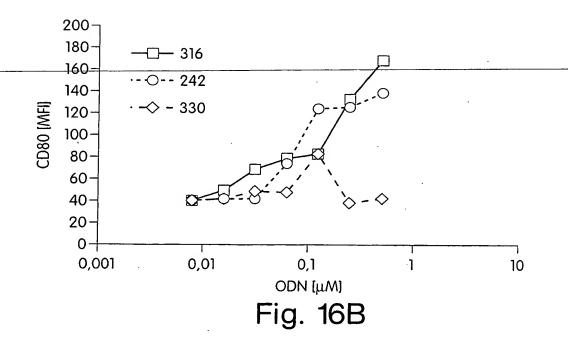


Fig. 14B



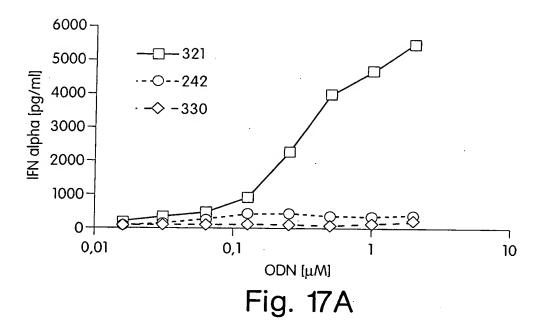


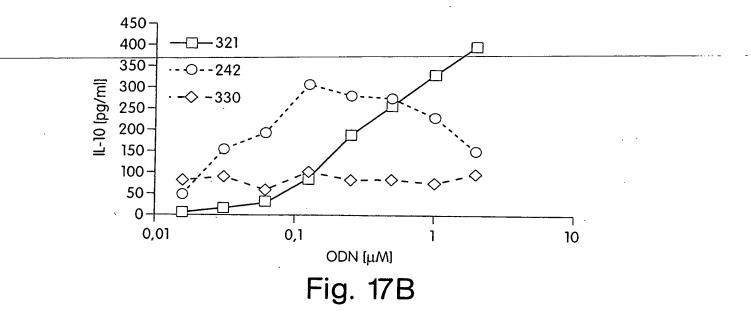




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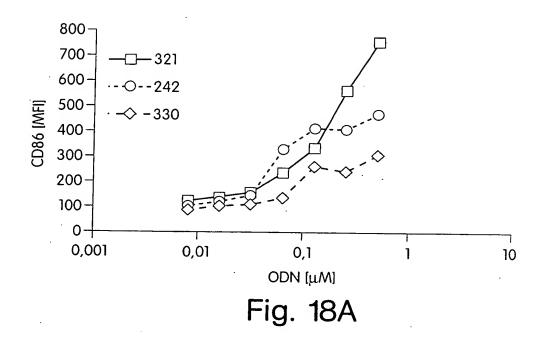
Docket No.: C01037.70048.US

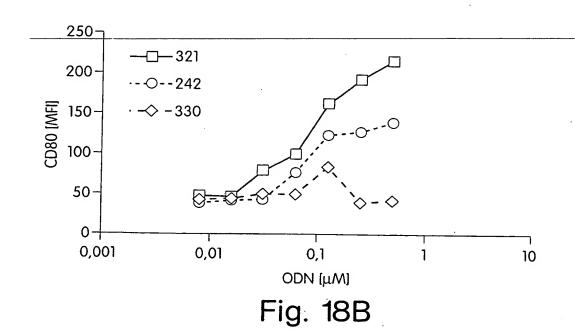


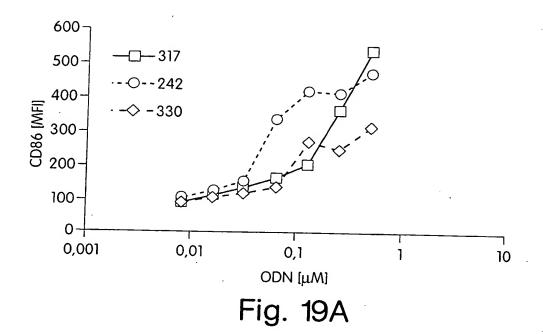


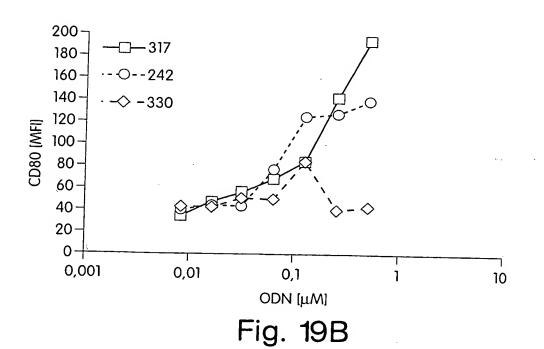
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Docket No.: C01037.70048.US

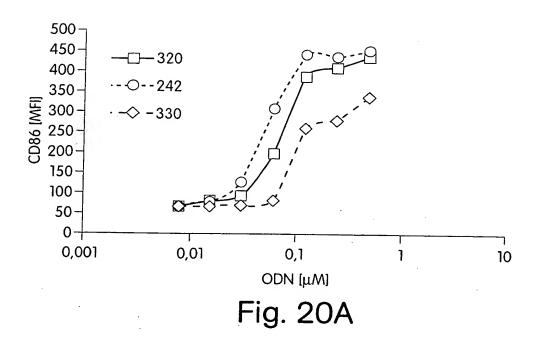








Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US



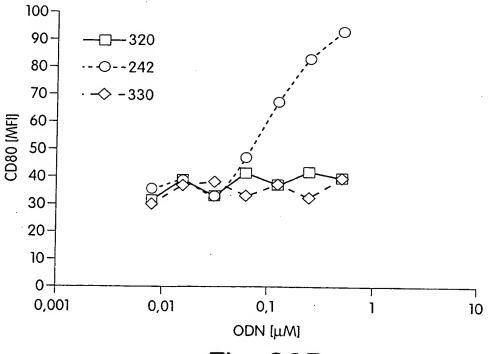
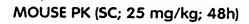


Fig. 20B

Fig. 21



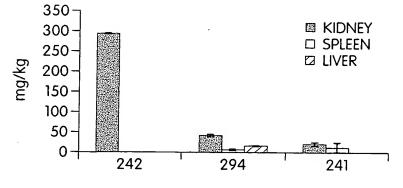


Fig. 22

Inventors: Arthur M. Krieg et al.

Title: IMMUNOSTIMULATORY NUCLEIC ACIDS Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US

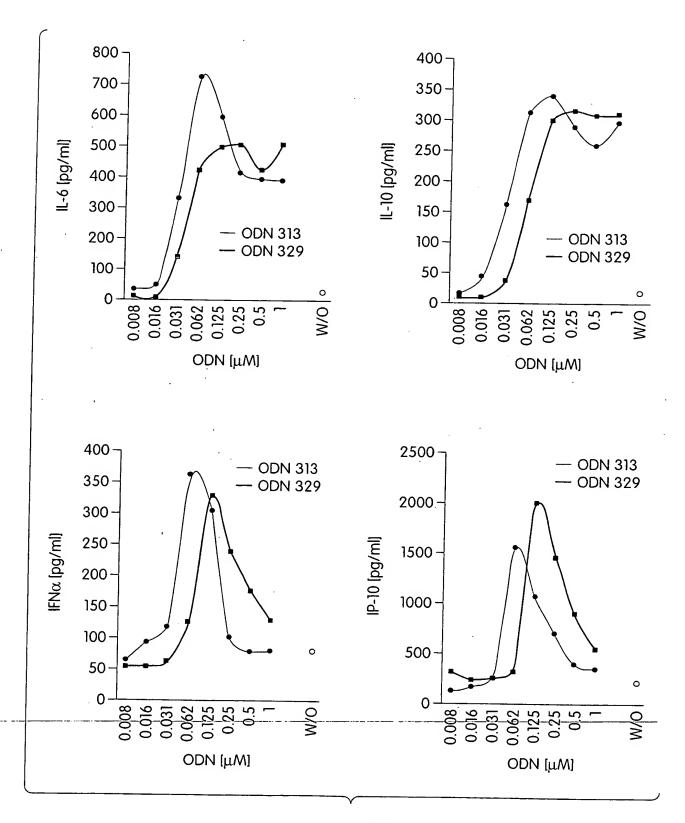


Fig. 23

Express Mail Label No.: EV292547294US Inventors: Arthur M. Krieg et al.

Title: IMMUNOSTIMULATORY NUCLEIC ACIDS

Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US

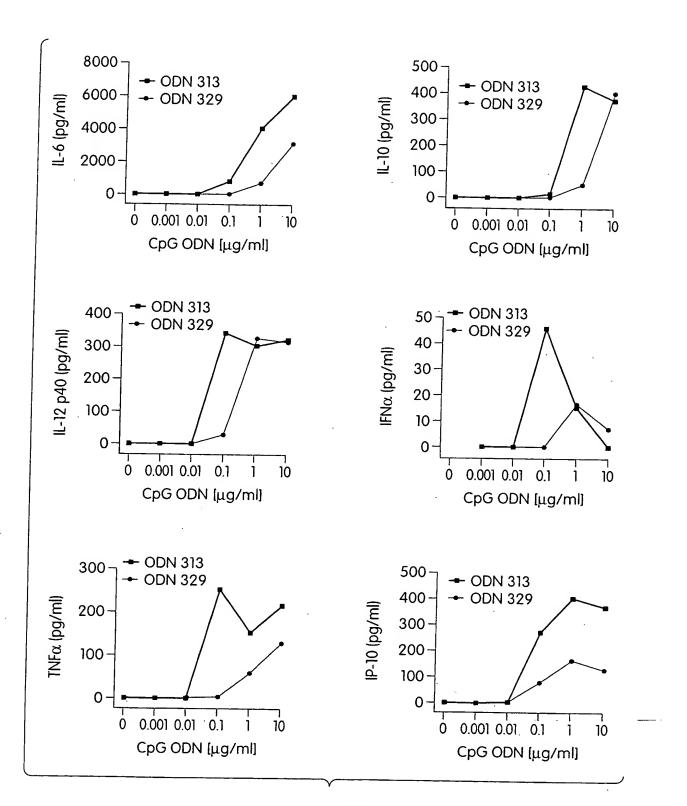
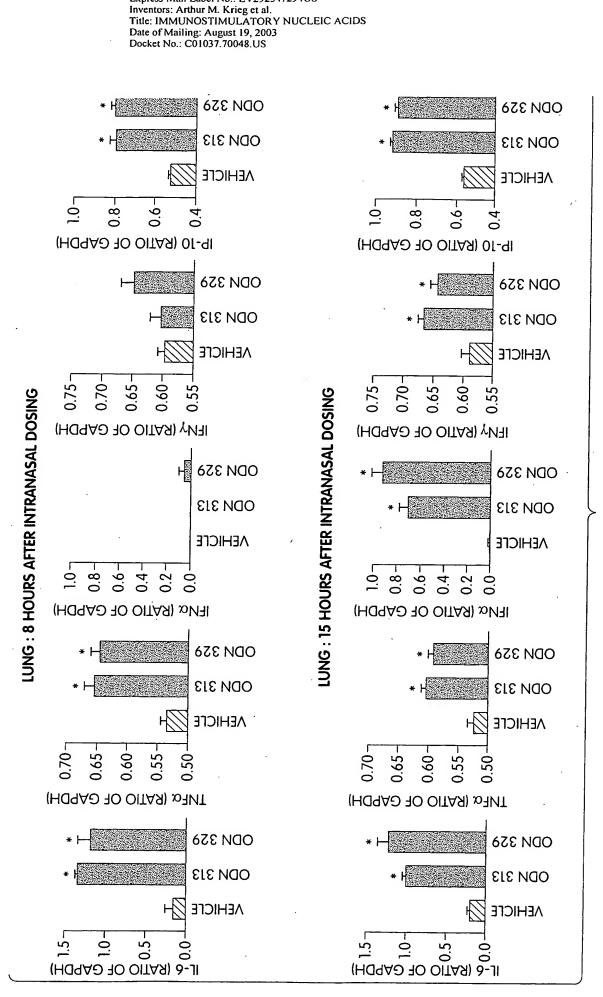


Fig. 24



IL-6 (RATIO OF GAPDH)

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Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US

IL-6 (RATIO OF GAPDH)

Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS

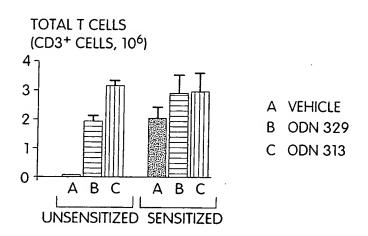
Fig. 25B-1

Inventors: Arthur M. Krieg et al. Title: IMMUNOSTIMULATORY NUCLEIC ACIDS Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US **ODN 356** ODN 313 **NEHICLE** <u>0</u>. 0.8 0.6 (HO9A5) OF-91 **ODN 356 ODN 313 NEHICLE** KIDNEY: 8 HOURS AFTER INTRAVENOUS DOSING 0.60 0.55 IFNY (RATIO OF GAPDH) **ODN 356** ODN 313 **NEHICLE** IFNa (RATIO OF GAPDH) **ODN 356 ODN 313 NEHICLE** 0.50 0.45 0.55 THF $\alpha$  (RATIO OF GAPDH) **ODN 356 ODN 313 NEHICLE** 

IL-6 (RATIO OF GAPDH)

Express Mail Label No.: EV292547294US

Fig. 25B-2



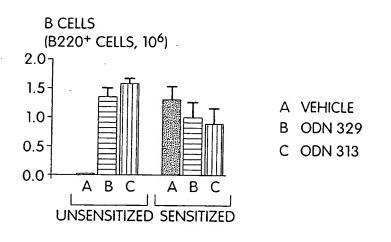


Fig. 26

Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US



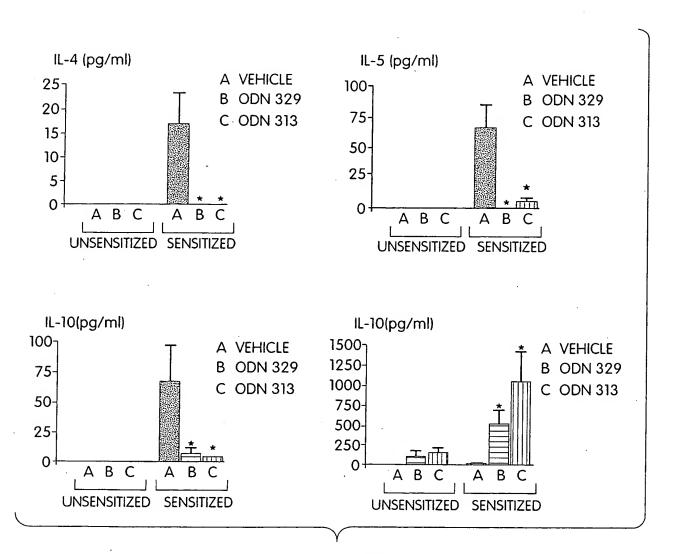
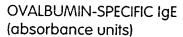
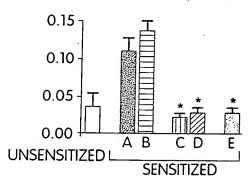


Fig. 27

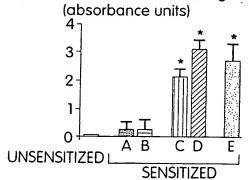
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Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US





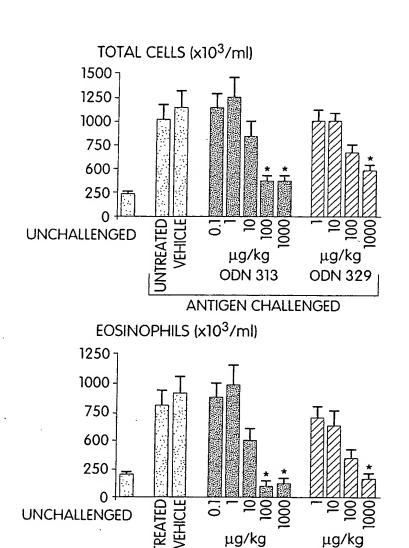
- A UNTREATED
- **B VEHICLE**
- C ODN 313 0.15mg/kg
- D ODN 313 1.5mg/kg
- E ODN 329 1.5mg/kg

## OVALBUMIN-SPECIFIC IgG2a



- A UNTREATED
- **B VEHICLE**
- C ODN 313 0.15mg/kg
- D ODN 313 1.5mg/kg
- E ODN 329 1.5mg/kg

Fig. 28



CD3+CD4+ (x10<sup>3</sup>/ml) 200 -150 100 50 0 🏗 UNCHALLENGED μg/kg **ODN 313** ANTIGEN CHALLENGED

**ODN 313** 

ANTIGEN CHALLENGED

**ODN 329** 

Fig. 29

Express Mail Label No.: EV292547294US
Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS

Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US

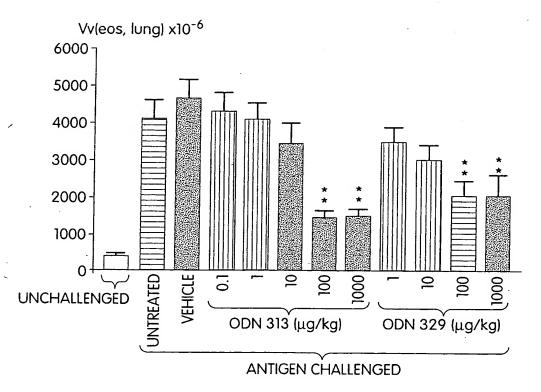


Fig. 30

Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US



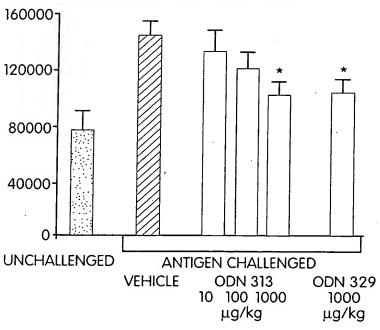
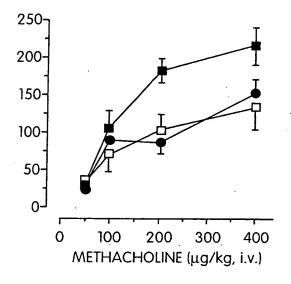


Fig. 31

Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US

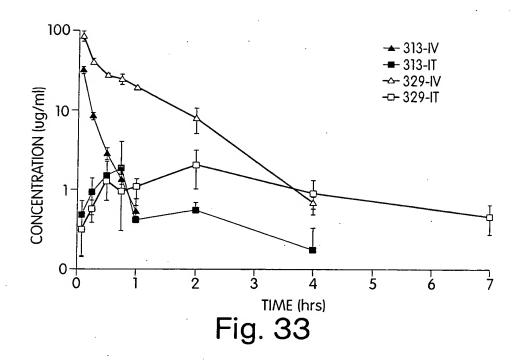
#### BRONCHOCONSTRICTION TO METHACHALOINE (% INCREASE IN AIRWAY RESISTANCE)

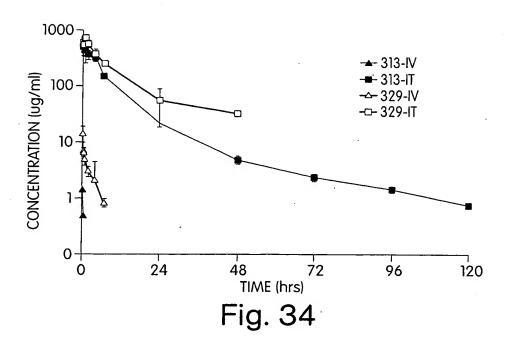


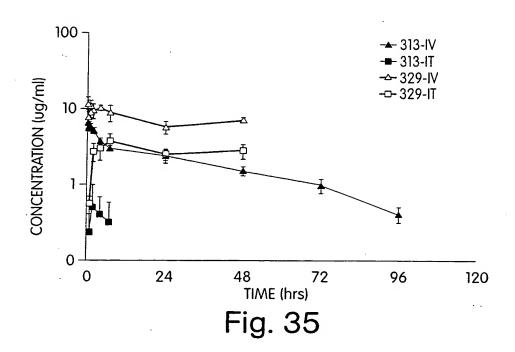
- UNCHALLENGED
- ANTIGEN CHALLENGED, TREATED WITH VEHICLE
- ANTIGEN CHALLENGED, TREATED WITH VEHICLE ODN 313 (1000 μg/kg)

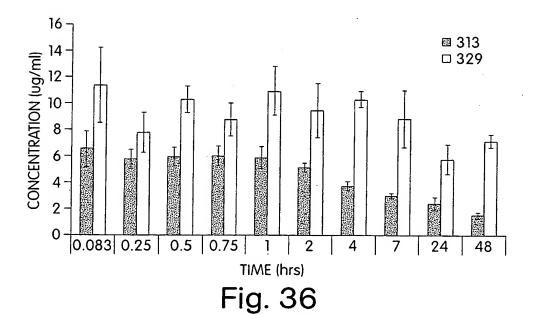
Fig. 32

Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US

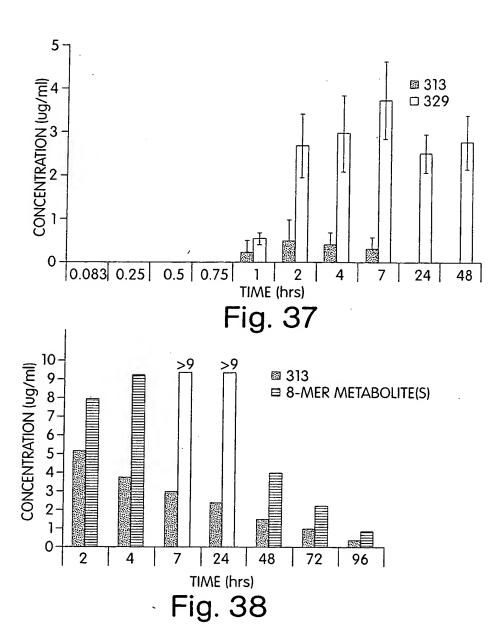


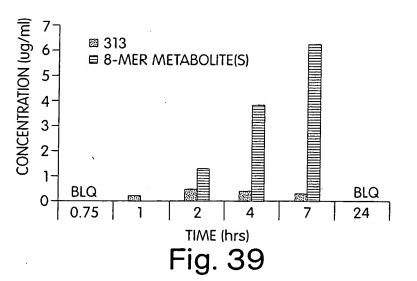




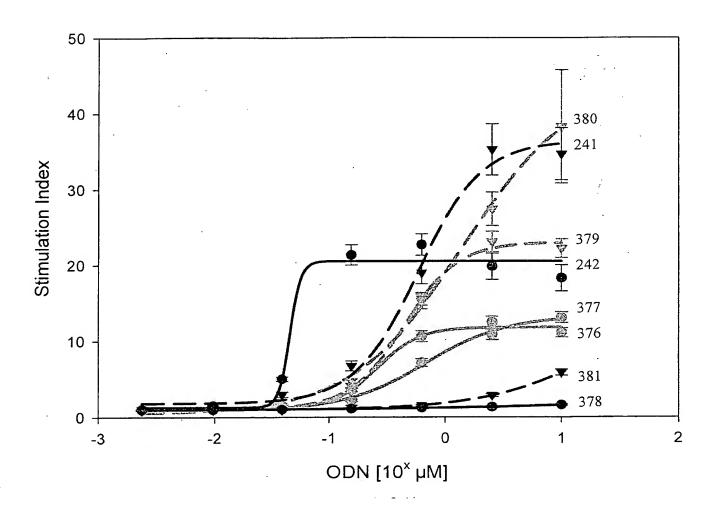


Date of Mailing: August 19, 2003 Docket No.: C01037.70048.US

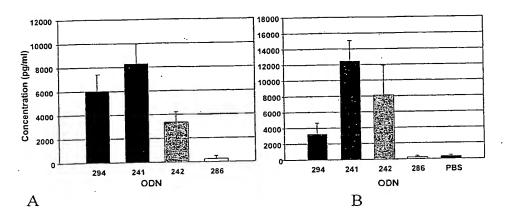




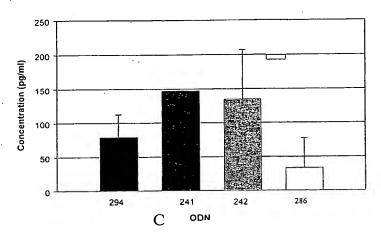
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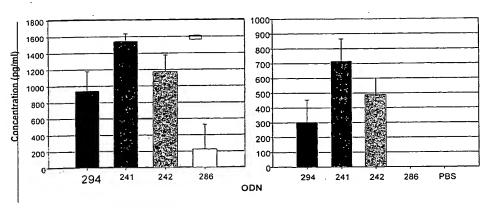
Inventors: Arthur M. Krieg et al.
Title: IMMUNOSTIMULATORY NUCLEIC ACIDS
Date of Mailing: August 19, 2003
Docket No.: C01037.70048.US



IFN-γ Induction (in plasma)



TNF- $\alpha$  Induction (in plasma)

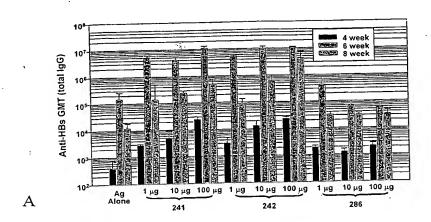


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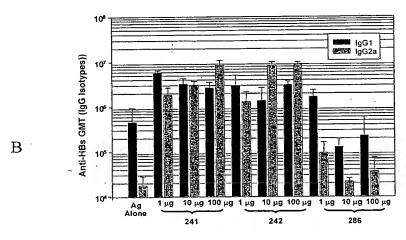
E

42

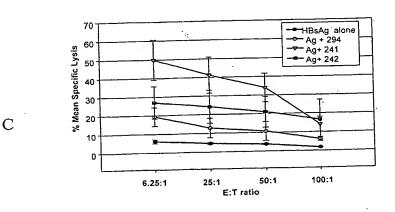
### Antibody Responses (10tal 190,



#### Nature of the antibody Response (IgG1 vs. IgG2a)



Cellular Responses (CTL)



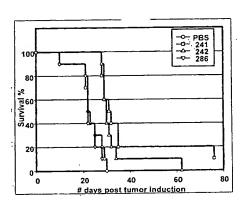
43

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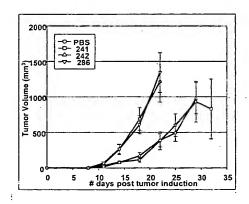
1200 (mm³) Day vs PBS
Day vs 242 1ug
Day vs 242 1oug
Day vs 242 3oug
Day vs 242 100ug **Tumor Volume** 600 50 # days post tumor induction

В



60

70



C

E

(mm<sub>3</sub>)

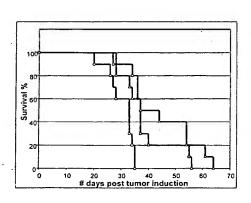
600

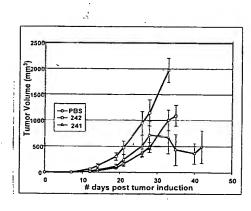
200

**Tumor Volume** 

A

30 40 50 60 # days post tumor induction





F

D

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